### 10/572932 PCT/JP2004/013722

# JAP9 Rec'd PCT/PTO 2 1 MAR 2006

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### SEQUENCE LISTING

<110> ONCOTHERAPY SCIENCE, INC.
THE UNIVERSITY OF TOKYO

<120> METHOD FOR DIAGNOSING HEPATOCELLULAR CARCINOMAS

<130> ONC-A0305P

<150> US 60/505, 632

<151> 2003-09-24

<160> 28

<170> PatentIn version 3.1

<210> 1

<211> 1528

<212> DNA

<213> Homo sapiens

<220>

<221> CDS

<222> (133).. (1308)

<223>

411

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ctc	gccci	tac	tgago	egago	g go	ccg	ggcg	g cce	gaggg	ggtc	cgce	gccg	ege g	gggg	egcaco	120
gcco	ctgg	ccg (	cc at	tg te	ge to	cc ca	ag ci	tc te	gg ti	tc ct	tg ac	g ga	ac cg	gg cg	gc ato	171
			Me	et Cy	rs Se	er G	ln Le	eu Ti	rp Pl	ne Le	eu Th	r As	sp Ai	rg Ai	rg Ile	)
			1				5					10	)			
											•					
cgc	gag	gac	tac	ccg	cag	gtg	cag	atc.	ctg	cgc	gcc	ctc	cgg	cag	cgc	219
Arg	Glu	Asp	Tyr	Pro	Gln	Val	Gln	Ile	Leu	Arg	Ala	Leu	Arg	Gln	Arg	
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tgc	tcc	gag	cag	gac	gtg	cgc	ttc	cgg	gcg	gtg	ctt	atg	gac	cag	atc	267
Cys	Ser	Glu	Gln	Asp	Val	Arg	Phe	Arg	Ala	Val	Leu	Met	Asp	Gln	Ile	
30					35				•	40					45	
gcc	gtc	acc	atc	gtc	ggc	ggc	cac	ctc	ggc	ctc	cag	cta	aac	cag	aag	319
Ala	Val	Thr	Ile	Val	Gly	Gly	His	Leu	Gly	Leu	Gln	Leu	Asn	Gln	Lys	
				50					55					60		
gcc	ctc	acc	act	ttc	ccg	gat	gtg	gtg	ctt	gta	cgg	gta	ccc	aca	ccc	368
Ala	Leu	Thr	Thr	Phe	Pro	Asp	Val	Val	Leu	Val	Arg	Val	Pro	Thr	Pro	
			65					70					75			

tca gtg cag tca gac agt gac atc act gtc ctg cga cac ctg gag aag

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Ser Val Gln Ser Asp Ser Asp Ile Thr Val Leu Arg His Leu Glu Lys 80 85 90

ctg ggc tgc cgg ttg gtc aat cgc cca cag agc atc tta aat tgc atc

Leu Gly Cys Arg Leu Val Asn Arg Pro Gln Ser Ile Leu Asn Cys Ile

95 100 105

aac aaa ttc tgg acg ttc caa gaa ctg gct gga cat ggg gtc ccc atg 507 Asn Lys Phe Trp Thr Phe Gln Glu Leu Ala Gly His Gly Val Pro Met 110 125

cca gac acc ttc tcc tat ggt ggg cat gaa gac ttt tca aaa atg att 555

Pro Asp Thr Phe Ser Tyr Gly Gly His Glu Asp Phe Ser Lys Met Ile

130 135 140

gat gaa gct gag ccc ctg ggc tac cca gtc gtg gtg aag agc aca cga 603
Asp Glu Ala Glu Pro Leu Gly Tyr Pro Val Val Val Lys Ser Thr Arg
145 150 155

ggc cac cgg gga aaa gct gtt ttt ctg gca aga gat aaa cat cac ctc 651 Gly His Arg Gly Lys Ala Val Phe Leu Ala Arg Asp Lys His His Leu 160 165 170

tct gac atc tgc cat ctg atc cgc cac gat gtg ccc tac ctg ttc cag

Ser Asp Ile Cys His Leu Ile Arg His Asp Val Pro Tyr Leu Phe Gln

175 180 185

1035

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aag	tac	gtg	aag	gag	tcc	cat	gga	aag	gac	atc	cgg	gtg	gtg	gtg	gta	747
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ggg	ggc	cag	gtc	ata	ggc	tct	atg	ctt	cgc	tgc	tcc	act	gat	gga	cgg	795
Gly	Gly	Gln	Val	Ile	Gly	Ser	Met	Leu	Arg	Cys	Ser	Thr	Asp	Gly	Arg	
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													tgt			843
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INT			GIY	Lys	GIN	Leu		116	GIN	vai	Ser		Ile	Leu	Gly	
		240					245					250				
ato	gac	ttc	tøt	ggr	att	σat	ctc	ctt	atc	ata	gar	as t	ggc	tee		939
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	255	1.110	0,0		110	260	Dou		110	nic t	265	пор	dly	DCI	·	
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gtg	gtg	tgt	gag	gca	aat	gct	aat	gtt	ggc	ttc	cta	gcc	ttt	gac	cag	987
	'							•					Phe			-
270					275					280				_	285	

gca tgc aac tta gat gtg ggt ggg atc att gca gac tat acc atg tcc

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Ala Cys Asn Leu Asp Val Gly Gly Ile Ile Ala Asp Tyr Thr Met Ser
290 295 300

ttg ctg cca aat agg cag act gga aag atg gct gtc ctc cca gga ctg 1083

Leu Leu Pro Asn Arg Gln Thr Gly Lys Met Ala Val Leu Pro Gly Leu

305 310 315

tcg agt cca agg gag aag aac gag ccg gat ggc tgt gct tca gct cag 1131
Ser Ser Pro Arg Glu Lys Asn Glu Pro Asp Gly Cys Ala Ser Ala Gln
320 325 330

gga gtt gca gag agc gtc tat acc atc aac agt ggg tct acc tct agc 1179

Gly Val Ala Glu Ser Val Tyr Thr Ile Asn Ser Gly Ser Thr Ser Ser

335 340 345

gaa agt gag cct gaa ctg gga gag atc cgg gat tcc tca gca agc aca 1227 Glu Ser Glu Pro Glu Leu Gly Glu Ile Arg Asp Ser Ser Ala Ser Thr 350 355 360 365

atg ggg gcc cca ccc tcc atg ctg ccc gaa cct ggc tac aac att aac 1275

Met Gly Ala Pro Pro Ser Met Leu Pro Glu Pro Gly Tyr Asn Ile Asn

370 375 380

aac agg att gct tct gag tta aaa ctt aag tga attcctgctt tttggcagca 1328 Asn Arg Ile Ala Ser Glu Leu Lys Leu Lys

385

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tttaaaccaa atcctactgc ttccctagta gttttgagtg aataaaatct ggactaatgt 1388
gatttcattt gcacagaaac tagaaatccc atctgggcac tcagcatttt ttctaacgat 1448
gatttaagca aatggcctag ctttgtggtt tttacaaaga caaatataaa aacactcaca 1508
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⟨210⟩ 2

⟨211⟩ 391

<212> PRT

<213≻ Homo sapiens

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Tyr Pro Gln Val Gln IIe Leu Arg Ala Leu Arg Gln Arg Cys Ser Glu 20 25 30

Gln Asp Val Arg Phe Arg Ala Val Leu Met Asp Gln Ile Ala Val Thr

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35

40

45

Ile Val Gly Gly His Leu Gly Leu Gln Leu Asn Gln Lys Ala Leu Thr
50 55 60

Thr Phe Pro Asp Val Val Leu Val Arg Val Pro Thr Pro Ser Val Gln 65 70 75 80

Ser Asp Ser Asp Ile Thr Val Leu Arg His Leu Glu Lys Leu Gly Cys
85 90 95

Arg Leu Val Asn Arg Pro Gln Ser Ile Leu Asn Cys Ile Asn Lys Phe
100 105 110

Trp Thr Phe Gln Glu Leu Ala Gly His Gly Val Pro Met Pro Asp Thr
115 120 125

Phe Ser Tyr Gly Gly His Glu Asp Phe Ser Lys Met Ile Asp Glu Ala 130 135 140

Glu Pro Leu Gly Tyr Pro Val Val Val Lys Ser Thr Arg Gly His Arg 145 150 155 160

Gly Lys Ala Val Phe Leu Ala Arg Asp Lys His His Leu Ser Asp Ile 165 170 175

Cys His Leu Ile Arg His Asp Val Pro Tyr Leu Phe Gln Lys Tyr Val 180 185 190

Lys Glu Ser His Gly Lys Asp Ile Arg Val Val Val Gly Gln
195 200 205

Val Ile Gly Ser Met Leu Arg Cys Ser Thr Asp Gly Arg Met Gln Ser 210 215 220

Asn Cys Ser Leu Gly Gly Val Gly Val Lys Cys Pro Leu Thr Glu Gln 225 230 235 240

Gly Lys Gln Leu Ala Ile Gln Val Ser Asn Ile Leu Gly Met Asp Phe

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245

250

255

Cys Gly Ile Asp Leu Leu Ile Met Asp Asp Gly Ser Phe Val Val Cys
260 265 270

Glu Ala Asn Ala Asn Val Gly Phe Leu Ala Phe Asp Gln Ala Cys Asn 275 280 285

Leu Asp Val Gly Gly Ile Ile Ala Asp Tyr Thr Met Ser Leu Leu Pro 290 295 300

Asn Arg Gln Thr Gly Lys Met Ala Val Leu Pro Gly Leu Ser Ser Pro 305 310 315 320

Arg Glu Lys Asn Glu Pro Asp Gly Cys Ala Ser Ala Gln Gly Val Ala
325
330
335

Glu Ser Val Tyr Thr Ile Asn Ser Gly Ser Thr Ser Ser Glu Ser Glu

340

345

350

Pro Glu Leu Gly Glu Ile Arg Asp Ser Ser Ala Ser Thr Met Gly Ala 355 360 365

Pro Pro Ser Met Leu Pro Glu Pro Gly Tyr Asn Ile Asn Asn Arg Ile 370 375 380

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<210> 3

<211> 22

<212> DNA

<213> Artificial

<220>

<223> An artificially synthesized primer sequence for RT-PCR

<400> 3

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22

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⟨211⟩ 20

<212> DNA

<213> Artificial

<220>

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ggtccaccac tgacacgttg

20

<210> 5

**⟨211⟩** 23

<212> DNA

<213> Artificial

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<223> An artificially synthesized primer sequence for RT-PCR

**<400>** 5

caaataggca gactggaaag atg

23

⟨210⟩ 6

<211> 23

<212> DNA

<213> Artificial

<220>

<223> An artificially synthesized primer sequence for RT-PCR

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23

⟨210⟩ 7

<211> 30

<212> DNA

<213≯ Artificial

<220>

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30

<210> 8

<211> 36

<212> DNA

<213> Artificial

<220>

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<400> 8

aatctcgaga gcaggaattc acttaagttt taactc

36

<210> 9

<211> 22

<212> DNA

<213> Artificial

<220>

<223> An artificially synthesized primer sequence for RT-PCR

**<400>** 9

tggtagccaa gtgcaggtta ta

22

<210> 10

<211> 22

<212> DNA

<213> Artificial

<220>

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<400> 10

ccaaagggtt tctgcagttt ca

22

<210> 11

<211> 30

<212> DNA

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<220>

<223> An artificially synthesized primer sequence for RT-PCR

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⟨210⟩ 12

<211> 29

<212> DNA

<213> Artificial

<220>

<223> An artificially synthesized primer sequence for RT-PCR

<400> 12

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29

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<211> 47

<212> DNA

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<220>

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**<400>** 13

47

<210> 14

**<211>** 34

<212> DNA

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<210> 15

<211> 51

<212> DNA

<213> Artificial

<220>

<223> An artificially synthesized oligonucleotide sequence for siRNA

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51

<210> 16

⟨211⟩ 51

<212> DNA

<213> Artificial

<220>

<223> An artificially synthesized oligonucleotide sequence for siRNA

<400> 16

aaaagaagca gcacgacttc ttctctcttg aagaagaagt cgtgctgctt c

51

**⟨211⟩** 51

<212> DNA

<213> Artificial

<220>

<223> An artificially synthesized oligonucleotide sequence for siRNA

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tcccgtgtcc gctgacagaa caattcaaga gattgttctg tcagcggaca c

51

⟨210⟩ 18

<211> 51

<212> DNA

<213> Artificial

<220>

<223> An artificially synthesized oligonucleotide sequence for siRNA

**<400>** 18

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51

<210> 19

<211> 19

<212> DNA

<213≯ Artificial

<220>

<223> An artificially synthesized target sequence for siRNA

<400> 19

gtgtccgctg acagaacaa

19

<210> 20

<211> 21

<212> DNA

<213≯ Artificial

<220>

<223> An artificially synthesized primer sequence for RT-PCR

<400> 20

gagctcctga accatctgct c

21

<210> 21

<211> 23

<212> DNA

<213> Artificial

<220>

<223> An artificially synthesized primer sequence for RT-PCR

<400> 21

caagatgtac agagcatcac agc

23

<210> 22

<211> 29

<212> DNA

<213> Artificial

<220>

 $\langle 223 \rangle$  An artificially synthesized primer sequence for RT-PCR

**<400>** 22

attgaattcg catggcgcca cccgcggcg

29

<210> 23

<211> 32

<212> DNA

<213> Artificial

<220>

<223> An artificially synthesized primer sequence for RT-PCR

**<400>** 23

aatggtacct caccaaggcc tccagacact cc

32

<210> 24

⟨211⟩ 51

<212> DNA

<213> Artificial

<220>

<223> An artificially synthesized oligonucleotide sequence for siRNA

**<400> 24** 

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51

<210> 25

<211> 51

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<213> Artificial

<220>

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**<400>** 25

21/27

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<210> 26

<211> 19

<212> DNA

<213> Artificial

<220>

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**<400>** 26

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19

<210> 27

<211> 1375

<212> DNA

<213> Homo sapiens

<220>

<221> CDS

**⟨222⟩** (125).. (799)

<223>

<400> 27

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gggc atg gcg cca ccc gcg gcg  Met Ala Pro Pro Ala Ala  1 5		
gat gag gac ggc tgg gag acg c Asp Glu Asp Gly Trp Glu Thr A		
ctg gtg gag aag aag cgg cgc g Leu Val Glu Lys Lys Arg Arg A 35		
ctg cgg ctg ctg ctg gcg ggc g Leu Arg Leu Leu Leu Ala Gly A 50 5		
gcc gaa gtg ctg gag ctg acg g Ala Glu Val Leu Glu Leu Thr V 65 70		
ggc cgg gcg cgc gag cgc gag c Gly Arg Ala Arg Glu Arg Glu G		

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80 85 90 95

ttc gct gcc ggc tac atc cag tgc atg cac gag gtg cac acg ttc gtg

457

Phe Ala Ala Gly Tyr Ile Gln Cys Met His Glu Val His Thr Phe Val

100

105

110

tcc acg tgc cag gcc atc gac gct acc gtc gct gcc gag ctc ctg aac 505

Ser Thr Cys Gln Ala Ile Asp Ala Thr Val Ala Ala Glu Leu Leu Asn
115 120 125

cat ctg ctc gag tcc atg ccg ctg cgt gag ggc agc agc ttc cag gat 553

His Leu Leu Glu Ser Met Pro Leu Arg Glu Gly Ser Ser Phe Gln Asp
130 135 140

ctg ctg ggg gac gcc ctg gcg ggg cca cct aga gcc cct gga cgg agt 601 Leu Leu Gly Asp Ala Leu Ala Gly Pro Pro Arg Ala Pro Gly Arg Ser 145 150 155

ggc tgg cct gcg ggg ggc gct ccg gga tcc cca ata ccc agc ccc ccg

Gly Trp Pro Ala Gly Gly Ala Pro Gly Ser Pro IIe Pro Ser Pro Pro

160 165 170 175

ggt cct ggg gac gac ctg tgc tcc gac ctg gag gag gcc cct gag gct

Gly Pro Gly Asp Asp Leu Cys Ser Asp Leu Glu Glu Ala Pro Glu Ala

180 185 190

### 24/27

gaa ctg agt cag gct cct gct gag ggg ccc ga	ac ttg gtg ccc gca gcc 745	į
Glu Leu Ser Gln Ala Pro Ala Glu Gly Pro As	sp Leu Val Pro Ala Ala	
195 200	205	
ctg ggc agc ctg acc aca gcc caa att gcc cg	gg agt gtc tgg agg cct 793	l I
Leu Gly Ser Leu Thr Thr Ala Gln Ile Ala Ai	g Ser Val Trp Arg Pro	
210 215	220	
tgg tga ccaatgccag ccagagtcct gcgggggtgg g	gcccggccct ccctggatct 849	)
Trp		
·		
cctcctcct cccaggggtt cagatgtggt ggggtaggg	gc cctggaagtc tcccaggtct 909	1
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acceageett caccageeet gtgegggete tgggggeag	ga ggtggcagga atggtgctgg 1149	)
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gtttgtagca cacttgagtt tgtgtattcc attgacatca aatgtgacaa ttttactaaa 1329

1375

<210> 28

<211> 224

<212> PRT

<213> Homo sapiens

**<400>** 28

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Val Glu Lys Lys Arg Arg Ala Arg Ile Asn Glu Ser Leu Gln Glu Leu 35 40 45

Arg Leu Leu Ala Gly Ala Glu Val Gln Ala Lys Leu Glu Asn Ala 50 . 55 60

Glu Val Leu Glu Leu Thr Val Arg Arg Val Gln Gly Val Leu Arg Gly
65 70 75 80

Arg Ala Arg Glu Arg Glu Gln Leu Gln Ala Glu Ala Ser Glu Arg Phe
85 90 95

Ala Ala Gly Tyr Ile Gln Cys Met His Glu Val His Thr Phe Val Ser 100 105 110

Thr Cys Gln Ala Ile Asp Ala Thr Val Ala Ala Glu Leu Leu Asn His
115 120 125

Leu Leu Glu Ser Met Pro Leu Arg Glu Gly Ser Ser Phe Gln Asp Leu 130 135 140

Leu Gly Asp Ala Leu Ala Gly Pro Pro Arg Ala Pro Gly Arg Ser Gly
145 150 155 160

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Trp Pro Ala Gly Gly Ala Pro Gly Ser Pro Ile Pro Ser Pro Pro Gly
165 170 175

Pro Gly Asp Asp Leu Cys Ser Asp Leu Glu Glu Ala Pro Glu Ala Glu
180 185 190

Leu Ser Gln Ala Pro Ala Glu Gly Pro Asp Leu Val Pro Ala Ala Leu 195 200 205

Gly Ser Leu Thr Thr Ala Gln Ile Ala Arg Ser Val Trp Arg Pro Trp 210 215 220